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PATENT

#44 31ta 4-22-02

IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

In re Application of: Jørgensen et al.

Serial No.: To be assigned

Confirmation No: To be assigned

Group Art Unit: To be assigned

Filed: August 13, 2001

Examiner: To be assigned

For: Method For Stable Chromosomal Multi-Copy Integration Of Genes

PRELIMINARY AMENDMENT

Commissioner for Patents Washington, DC 20231

Sir:

1)

Before the above-captioned application is taken up for examination, entry of the following amendment is respectfully requested:

IN THE CLAIMS:

Please cancel claims 6-29, 45-62 and 64-65 without prejudice or disclaimer.

Please substitute the following amended claims for the pending claims having the same claim numbers:

O,

3. The method of claim 1, wherein subsequent to the step of introducing the DNA construct and cultivating the cell under selective conditions, or subsequent to the step of selecting a host cell, a second recombination takes place between the DNA fragment and the homologous host cell DNA sequence.



44. A host cell comprising at least two copies of a gene of interest stably integrated into the chromosome, where at least one copy is integrated adjacent to a conditionally essential *locus* and wherein the cell is obtainable by the method of claim 1.



66. A process for producing an enzyme of interest, comprising cultivating a cell of claim 44 under conditions appropriate for producing the enzyme, and optionally purifying the enzyme.